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(74) Agents: WALES, Michele, M. et al.; Human Genome Sciences, Inc., 9410 Key West Avenue, Rockville, MD 20850 (US). (81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

#### Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(54) Title: HUMAN PANCREAS AND PANCREATIC CANCER ASSOCIATED GENE SEQUENCES AND POLYPEPTIDES

#### (57) Abstract

This invention relates to newly identified pancreas or pancreatic cancer related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "pancreatic cancer antigens", and to the complete gene sequences associated therewith and to the expression products thereof, as well as the use of such pancreatic cancer antigens for detection, prevention and treatment of disorders of the pancreas, particularly the presence of pancreatic cancer. This invention relates to the pancreatic cancer antigens as well as vectors, host cells, antibodies directed to pancreatic cancer antigens and recombinant and synthetic methods for producing the same. Also provided are diagnostic methods for diagnosing and treating, preventing and/or prognosing disorders related to the pancreas, including pancreatic cancer, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of pancreatic cancer antigens of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and/or function of the polypeptides of the present invention.

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### What Is Claimed Is:

- 1. An isolated nucleic acid molecule comprising a polynucleotide having a nucleotide sequence at least 95% identical to a sequence selected from the group consisting of:
- (a) a polynucleotide fragment of SEQ ID NO:X or a polynucleotide fragment of the cDNA sequence included in the related cDNA clone, which is hybridizable to SEQ ID NO:X;
- (b) a polynucleotide encoding a polypeptide fragment of SEQ ID NO:Y or a polypeptide fragment encoded by the cDNA sequence included in the related cDNA clone, which is hybridizable to SEQ ID NO:X;
  - (c) a polynucleotide encoding a polypeptide fragment of a polypeptide encoded by SEQ ID NO:X or a polypeptide fragment encoded by the cDNA sequence included in the related cDNA clone, which is hybridizable to SEQ ID NO:X;
  - (d) a polynucleotide encoding a polypeptide domain of SEQ ID NO:Y or a polypeptide domain encoded by the cDNA sequence included in the related cDNA clone, which is hybridizable to SEQ ID NO:X;
  - (e) a polynucleotide encoding a polypeptide epitope of SEQ ID NO:Y or a polypeptide epitope encoded by the cDNA sequence included in the related cDNA clone, which is hybridizable to SEQ ID NO:X;
  - (f) a polynucleotide encoding a polypeptide of SEQ ID NO:Y or the cDNA sequence included in the related cDNA clone, which is hybridizable to SEQ ID NO:X, having biological activity;
    - (g) a polynucleotide which is a variant of SEQ ID NO:X;
    - (h) a polynucleotide which is an allelic variant of SEQ ID NO:X;
  - (i) a polynucleotide which encodes a species homologue of the SEQ ID NO:Y;
- (j) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(i), wherein said polynucleotide does not hybridize under stringent conditions to a nucleic acid molecule having a nucleotide

- 10. The recombinant host cell of claim 9 comprising vector sequences.
- 11. An isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence selected from the group consisting of:
  - (a) a polypeptide fragment of SEQ ID NO:Y or of the sequence encoded by the cDNA included in the related cDNA clone;
  - (b) a polypeptide fragment of SEQ ID NO:Y or of the sequence encoded by the cDNA included in the related cDNA clone, having biological activity;
- (c) a polypeptide domain of SEQ ID NO:Y or of the sequence encoded by the cDNA included in the related cDNA clone;
  - (d) a polypeptide epitope of SEQ ID NO:Y or of the sequence encoded by the cDNA included in the related cDNA clone;
- (e) a full length protein of SEQ ID NO:Y or of the sequence encoded by the cDNA included in the related cDNA clone;
  - (f) a variant of SEQ ID NO:Y;
  - (g) an allelic variant of SEQ ID NO:Y; or
  - (h) a species homologue of the SEQ ID NO:Y.
- 20 12. The isolated polypeptide of claim 11, wherein the full length protein comprises sequential amino acid deletions from either the C-terminus or the N-terminus.
- 13. An isolated antibody that binds specifically to the isolated polypeptide 25 of claim 11.
  - 14. A recombinant host cell that expresses the isolated polypeptide of claim 11.
- 30 15. A method of making an isolated polypeptide comprising:

- 21. The gene corresponding to the cDNA sequence of SEQ ID NO:Y.
- 22. A method of identifying an activity in a biological assay, wherein the method comprises:
- 5 (a) expressing SEQ ID NO:X in a cell;
  - (b) isolating the supernatant;
  - (c) detecting an activity in a biological assay; and
  - (d) identifying the protein in the supernatant having the activity.
- The product produced by the method of claim 20.

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N15/11 C12N15/12

C12N15/11 C12N15/12 C07K14/47 C07K16/18 G06F17/30 C12N15/62 A61K31/713 C12N15/86 C12Q1/68 C12N5/10 G01N33/577

According to International Patent Classification (IPC) or to both national classification and IPC

#### **B. FIELDS SEARCHED**

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, MEDLINE, WPI Data

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	DATABASE EMBL [Online] 26 December 1999 (1999-12-26) BIRREN B. ET AL: "Homo sapiens chromosome 11, clone RP11-707D17" Database accession no. AC18997 XP002170377 nts. 20-1487	2-4, 15-17, 21-26
P,X	WO 00 55320 A (HUMAN GENOME SCIENCES INC; ROSEN CRAIG A (US); RUBEN STEVEN M (US)) 21 September 2000 (2000-09-21) SEQ ID NOs.171 and 630 page 494; claim 19	2-4, 15-27, 29-33
A	WO 93 16180 A (MAX PLANCK GESELLSCHAFT; UNI DEGLI STUDI G D ANNUNZIO C (IT)) 19 August 1993 (1993-08-19)	

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
Special categories of cited documents:  "A" document defining the general state of the art which is not considered to be of particular relevance  "E" earlier document but published on or after the international filing date  "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  "O" document referring to an oral disclosure, use, exhibition or other means  "P" document published prior to the international filing date but later than the priority date claimed  Date of the actual completion of the international search	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.  "&" document member of the same patent family  Date of mailing of the international search report
Name and mailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (+31-70) 340-3016	Authorized officer  Blanco Urgoiti, B

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C (Continue	No. ) DOCUMENTS CONSIDERED TO DE TO	PC1/05 6	71/00099 .
C.(Continua Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages		Dolougit
J.,	, which appropriate, of the relevant passages		Relevant to claim No.
A	ANISOWICZ A ET AL: "A NOVEL PROTEASE HOMOLOG DIFFERENTIALLY EXPRESSED IN BREAST AND OVARIAN CANCER" MOLECULAR MEDICINE, US, BLACKWELL SCIENCE, CAMBRIDGE, MA, vol. 2, no. 5, 1 September 1996 (1996-09-01), pages 624-636, XP002060076 ISSN: 1076-1551		
A	BERGSTROM DAVID E ET AL: "Regulatory Autonomy and Molecular Characterization of the Drosophila out at first Gene." GENETICS, vol. 139, no. 3, 1995, pages 1331-1346, XP001009906 ISSN: 0016-6731 cited in the application		
	!		

## Information on patent family members

International Application No PCT/US 01/00899

Patent document cited in search report		Publication date	Patent family member(s)		Publication date
WO 0055320	A	21-09-2000	AU AU AU AU WO WO WO WO WO	3395900 A 3617600 A 3617700 A 3619400 A 3619500 A 3869400 A 0055173 A 0055350 A 0055351 A 0055180 A 0055174 A	04-10-2000 04-10-2000 04-10-2000 04-10-2000 04-10-2000 04-10-2000 21-09-2000 21-09-2000 21-09-2000 21-09-2000
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